WHAT IS CLAIMED IS:

5

- 1. Human aggrecanase present in other than its natural environment.
- 2. The aggrecanase according to Claim 1, wherein said aggrecanase has an amino acid sequence substantially identical to the sequence of SEQ ID NO:01.
 - 3. A fragment of the aggrecanase according to Claim 1.
- 4. A nucleic acid present in other than its natural environment, wherein said nucleic acid has
 10 a nucleotide sequence encoding aggrecanase.
 - 5. A nucleic acid according to Claim 4, wherein said nucleic acid has a nucleic acid sequence that is substantially identical to the nucleotide sequence of SEQ ID NO:02.
- 15 6. A fragment of the nucleic acid according to Claim 4.
 - 7. An isolated nucleic acid or mimetic thereof that hybridizes under stringent conditions to the nucleic acid according to Claim 4 or its complementary sequence.
- 8. An expression cassette comprising a transcriptional initiation region functional in an expression host, a nucleic acid having a nucleotide sequence found in the nucleic acid according to Claim 4 under the transcriptional regulation of said transcriptional initiation region, and a transcriptional termination region functional in said expression host.
- 9. A cell comprising an expression cassette according to Claim 8 as part of an extrachromosomal element or integrated into the genome of a host cell as a result of introduction of said expression cassette into said host cell.
 - 10. The cellular progeny of the host cell according to Claim 9.

30

- 11. A method of producing aggrecanase, said method comprising:
 growing a cell according to Claim 9, whereby said aggrecanase is expressed; and
 isolating said aggrecanase substantially free of other proteins.
- 5 12. A monoclonal antibody binding specifically to aggrecanase.
 - 13. The antibody according to Claim 12, wherein said antibody inhibits aggrecanase activity.
- 14. The monoclonal antibody according to Claim 13, wherein said antibody is selected from the group consisting of a human antibody or a humanized antibody.
 - 15. A method for modulating aggrecanase in a host, said method comprising:

 administering an effective amount of an aggrecanase modulatory agent to said host.
- 15 16. The method according to Claim 15, wherein said modulatory agent is a small molecule.
 - 17. The method according to Claim 15, wherein said modulatory agent is an antibody.
 - 18. The method according to Claim 15, wherein said modulatory agent is a nucleic acid.
 - 19. A method of screening to identify aggrecanase modulatory agents, said method comprising:

contacting aggrecanase with an aggrecanase substrate in the presence of an potential modulatory agent; and

- determining the effect of said modulatory agent on the activity of said aggrecanase.
 - 20. The method according to Claim 19, wherein said aggrecanase substrate comprises a gluala bond.

20

- 21. The method according to claim 20, wherein said aggrecanase substrate is aggrecan or a fragment thereof.
- A method of treating a host suffering from a disease condition associated with aggrecanase
 activity, said method comprising:
 administering to said host an aggrecanase modulatory agent.
 - 23. The method according to Claim 22, wherein said aggrecanase modulatory agent is an aggrecanase antagonist.
 - 24. The method according to Claim 23, wherein said disease condition is characterized by the presence of aggrecan cleavage products.
 - 25. A non-human transgenic animal model capable of expressing aggrecanase.

10